

### **ABSTRACT OF THE DISCLOSURE**

Human interleukin-18 (IL-18) polypeptides and substitution mutants thereof were conjugated to water-soluble polymers at specific sites on the human IL-18 protein. These  
5 conjugated human IL-18 and substitution mutants thereof retain biological activity. These conjugated cytokines demonstrate enhanced and unexpected biological properties when compared to the corresponding unconjugated cytokines.

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